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2 9. The method of claim 1, wherein expression of the candidate genes is inhibited by at least about 50%.

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2 10. The method of claim 1, wherein expression of the candidate genes is activated by at least about 150%.

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2 11. The method of claim 1, wherein the zinc finger proteins are fusion proteins comprising a regulatory domain.

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2 12. The method of claim 1, wherein expression of the zinc finger proteins is induced by administration of an exogenous agent.

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2 13. The method of claim 11, wherein the zinc finger proteins are fusion proteins comprising at least two regulatory domains.

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2 14. The method of claim 1, wherein the cell is selected from the group consisting of animal cell, a plant cell, a bacterial cell, a protozoal cell, or a fungal cell.

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2 15. The method of claim 14, wherein the cell is a mammalian cell

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2 16. The method of claim 15, wherein the cell is a human cell

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2 17. The method of claim 1, wherein the modulation of expression is activation of gene expression that prevents repression of gene expression.

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2 18. The method of claim 1, wherein the modulation of expression is inhibition of gene expression that prevents gene activation.

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2 19. The method of claim 11, wherein the regulatory domain is selected from the group consisting of a transcriptional repressor, a methyl transferase, a transcriptional activator, a histone acetyltransferase, and a histone deacetylase.

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2 20. The method of claim 1, wherein the first and second zinc finger proteins are encoded by an expression vector comprising a zinc finger protein nucleic acid operably linked to a promoter, and wherein the method further comprises the step of first administering the expression vector to the cell.

1 21. The method of claim 20, wherein expression of the zinc finger
2 proteins is under small molecule control.

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1 22. The method of claim 21, wherein expression of the first zinc finger
2 protein and expression of the second zinc finger protein are under different small
3 molecule control, wherein both the first and the second zinc finger protein are fusion
4 proteins comprising a regulatory domain, and wherein the first and the second zinc finger
5 proteins are expressed in the same cell.

1 23. The method of claim 22, wherein both the first and the second zinc
2 finger proteins comprise a regulatory domain that represses gene expression.

1 24. The method of claim 20, wherein the expression vector is a viral
2 vector.

1 25. The method of claim 24, wherein the expression vector is a
2 retroviral expression vector, an adenoviral expression vector, or an AAV expression
3 vector.

1 26. The method of claim 20, wherein the zinc finger proteins are
2 encoded by a nucleic acid operably linked to an inducible promoter.

1 27. The method of claim 1, wherein the cell comprises less than about
2 1.5×10^6 copies of each zinc finger protein.

1 28. The method of claim 1, wherein the target site is upstream of a
2 transcription initiation site of the candidate gene.

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1 29. The method of claim 1, wherein the target site is adjacent to a
2 transcription initiation site of the candidate gene.

1 30. The method of claim 1, wherein the target site is adjacent to an
2 RNA polymerase pause site downstream of a transcription initiation site of the candidate
3 gene.

1 31. A method of identifying the biological function of a candidate
2 gene, the method comprising the steps of:

- 3 (i) identifying a plurality of candidate genes;
 4 (ii) providing a first zinc finger protein that binds to a first target site of a
 5 first candidate gene;
 6 (iii) culturing a first cell under conditions where the first zinc finger
 7 protein contacts the first candidate gene, wherein the first zinc finger protein modulates
 8 expression of the first candidate gene;
 9 (iv) determining the expression pattern of the candidate genes and
 10 determining whether or not the first candidate gene is associated with the selected
 11 phenotype; and
 12 (v) repeating steps (ii)-(iv) for each candidate gene.

1 32. The method of claim 31, further comprising providing a second
 2 zinc finger protein that binds to a second target site of the first candidate gene.

1 33. The method of claim 31, wherein at least one of the candidate
 2 genes is an EST of at least about 200 nucleotides in length.

1 34. The method of claim 31, wherein at least two candidate genes are
 2 required to cause the selected phenotype.

1 35. The method of claim 31, wherein the candidate genes are
 2 endogenous genes.

1 36. The method of claim 31, wherein expression of the candidate genes
 2 is inhibited by at least about 50%.

1 37. The method of claim 31, wherein expression of the candidate genes
 2 is activated to at least about 150%.

1 38. The method of claim 31, wherein the zinc finger protein is a fusion
 2 protein comprising a regulatory domain.

1 39. The method of claim 38, wherein the regulatory domain is under
 2 small molecule control.

1 40. The method of claim 38, wherein the zinc finger proteins are fusion
 2 proteins comprising at least two regulatory domains.

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1 41. The method of claim 31, wherein the cell is selected from the
2 group consisting of animal cell, a plant cell, a bacterial cell, a protozoal cell, or a fungal
3 cell.

1 42. The method of claim 41, wherein the cell is a mammalian cell

1 43. The method of claim 42, wherein the cell is a human cell

1 44. The method of claim 31, wherein the modulation of expression is
2 activation of gene expression that prevents repression of gene expression.

1 45. The method of claim 31, wherein the modulation of expression is
2 inhibition of gene expression that prevents gene activation.

1 46. The method of claim 38, wherein the regulatory domain is selected
2 from the group consisting of a transcriptional repressor, a methyl transferase, a
3 transcriptional activator, a histone acetyltransferase, and a histone deacetylase.

1 47. The method of claim 31, wherein the zinc finger protein is encoded
2 by an expression vector comprising a zinc finger protein nucleic acid operably linked to a
3 promoter, and wherein the method further comprises the step of first administering the
4 expression vector to the cell.

1 48. The method of claim 47, wherein expression of the zinc finger
2 protein is under small molecule control.

1 49. The method of claim 47, wherein the expression vector is a viral
2 vector.

1 50. The method of claim 49, wherein the expression vector is a
2 retroviral expression vector, an adenoviral expression vector, or an AAV expression
3 vector.

1 51. The method of claim 47, wherein the zinc finger protein is encoded
2 by a nucleic acid operably linked to an inducible promoter.

1 52. The method of claim 31, wherein the cell comprises less than about
2 1.5×10^6 copies of the zinc finger protein.

1 53. The method of claim 31, wherein the target site is upstream of a
2 transcription initiation site of the candidate gene.

1 54. The method of claim 31, wherein the target site is adjacent to a
2 transcription initiation site of the candidate gene.

1 55. The method of claim 31, wherein the target site is adjacent to an
2 RNA polymerase pause site downstream of a transcription initiation site of the candidate
3 gene.

1 56. A method of identifying the biological function of a candidate
2 gene, the method comprising the steps of:

3 (i) selecting a first candidate gene;

4 (ii) providing a first zinc finger that binds to a first target site of the first
5 candidate gene and a second zinc finger that binds to a second target site of the first
6 candidate gene;

7 (iii) culturing a first cell under conditions where the first zinc finger
8 protein contacts the first candidate gene, and culturing a second cell under conditions
9 where the second zinc finger protein contacts the first candidate gene, wherein the first
10 and the second zinc finger proteins modulate expression of the first candidate gene; and

11 (iv) assaying for a selected phenotype, thereby identifying whether or not
12 the first candidate gene is associated with the selected phenotype.

1 57. The method of claim 56, further comprising providing a third zinc
2 finger protein that binds to a target site of a second candidate gene.

1 58. The method of claim 56, further comprising selecting a plurality of
2 candidate genes and providing a plurality of zinc finger proteins that bind to a target site
3 of each candidate gene.

1 59. The method of claim 57, wherein the second candidate gene is a
2 control gene.

1 60. The method of claim 56, wherein the first candidate gene is an EST
2 of at least about 200 nucleotides in length.

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75. The method of claim 56, wherein the first and the second zinc finger proteins are encoded by an expression vector comprising a zinc finger protein nucleic acid operably linked to a promoter, and wherein the method further comprises the step of first administering the expression vector to the cell.

2 proteins is under small molecule control, wherein both the first and the second zinc finger
1 77. The method of claim 76, wherein expression of the first zinc finger
2 protein and expression of the second zinc finger protein are under different small
3 molecule control, wherein both the first and the second zinc finger protein are fusion
4 proteins comprising a regulatory domain, and wherein the first and the second zinc finger
5 proteins are expressed in the same cell.

3 protein comprises a region of the protein that is located within the viral capsid.
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2 79. The method of claim 75, wherein the expression vector is a viral vector.

81. The method of claim 75, wherein the zinc finger proteins are encoded by a nucleic acid operably linked to an inducible promoter.

83. The method of claim 56, wherein the first or the second target site is upstream of a transcription initiation site of the first candidate gene.

1 85. The method of claim 56, wherein the first or the second target site
2 is adjacent to an RNA polymerase pause site downstream of a transcription initiation site
3 of the first candidate gene.

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